Appl. No. 10/684,796 Amdt. dated January 14, 2008 Reply to Office Action of July 13, 2007

## **Amendments to the Claims:**

This listing of claims will replace all prior versions, and listings of claims in the application:

## **Listing of Claims:**

- 1. (Currently Amended) A method of detecting PDZ polypeptide binding to an alpha adrenergic receptor, comprising
- a) combining a labeled-polypeptide containing an alpha <u>2</u> adrenergic receptor C-terminal peptide sequence <u>comprising the last 3 consecutive amino acids at the C-terminal end of SEQ ID NO: 26, SEQ ID NO: 27 or SEQ ID NO: 28 with a PDZ polypeptide *in vitro*, and</u>
- b) detecting <u>specific</u> binding between the PDZ polypeptide and the alpha <u>2</u> adrenergic receptor polypeptide.
- 2. (Currently Amended) The method of claim 1 wherein the <u>alpha 2</u> adrenergic receptor C-terminal peptide PL polypeptide is a biotinylated peptide.
- 3. (Currently Amended) The method of claim 1 wherein the <u>alpha 2</u> adrenergic receptor C-terminal peptide PL polypeptide is a fluorescence labeled peptide.
- 4. (Currently Amended) The method of claim 1 wherein the <u>alpha 2</u> adrenergic receptor C-terminal peptide PL polypeptide an epitope tagged is a protein expressed in a host cell.
- 5. **(Withdrawn)** A method of determining whether a test compound is a modulator of binding between a PDZ polypeptide and an alpha adrenergic PL polypeptide, comprising:
- (a) contacting under suitable binding conditions (i) a PDZ polypeptide, and (ii) a PL peptide, wherein

the PL peptide comprises a C-terminal sequence of the PL polypeptide,

Appl. No. 10/684,796 Amdt. dated January 14, 2008 Reply to Office Action of July 13, 2007

the PDZ polypeptide and the PL peptide are a binding pair as specified in Table 8; and

contacting is performed in the presence of the test compound; and

- (b) detecting formation of a complex between the PDZ-domain polypeptide and the PL peptide, wherein
- (i) presence of the complex at a level that is statistically significantly higher in the presence of the test compound than in the absence of test compound is an indication that the test compound is an agonist, and
- (ii) presence of the complex at a level that is statistically significantly lower in the presence of the test compound than in the absence of test compound is an indication that the test compound is an antagonist.
  - 6. (Withdrawn) The method of claim 5, wherein the modulator is a peptide.
- 7. **(Withdrawn)** A modulator of binding between a specific PDZ polypeptide and an alpha adrenergic receptor PL polypeptide, wherein the modulator is
- (a) a peptide comprising at least 3 residues of a C-terminal sequence demonstrated to bind the target PDZ polypeptide; or
  - (b) a peptide mimetic of the peptide of section (a); or
- (c) a small molecule having similar functional activity as the peptide of section (a) with respect to the PDZ polypeptide and PL polypeptide binding pair.
- 8. **(Withdrawn)** The modulator of claim 7 that modulates a specific interaction listed in Table 8.
  - 9. (Withdrawn) The modulator of claim 7 that is an agonist.
  - 10. (Withdrawn) The modulator of claim 7 that is an antagonist.

Appl. No. 10/684,796 Amdt. dated January 14, 2008 Reply to Office Action of July 13, 2007 **PATENT** 

- 11. **(Withdrawn)** A pharmaceutical composition comprising a modulator of claim 7.
- 12. **(Withdrawn)** A method of treating a disorder from Table 9, comprising administering a therapeutically effective amount of a modulator of claim 7, wherein the PDZ polypeptide and the alpha adrenergic receptor PL polypeptide are a binding pair as specified in Table 8.